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NOTICE OF ALLOWANCE AND FEE(S) DUE

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NIXON & VANDERHYE, PC 901 NORTH GLEBE ROAD, 11TH FLOOR ARLINGTON, VA 22203

| EXAMINER | | | | |
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| TON, THAIAN N | | | | |
| ART UNIT PAPER NUMBER | | | | |

1632

DATE MAILED: 03/10/2008

| APPLICATION NO. | FILING DATE | FIRST NAMED INVENTOR | ATTORNEY DOCKET NO. | CONFIRMATION NO. |
|-----------------|-------------|----------------------|---------------------|------------------|
| 10/502,332 | 07/23/2004 | Marie Malissen | BJS-3665-113 | 9194 |

TITLE OF INVENTION: MUTATED GENE CODING FOR A LAT PROTEIN AND THE BIOLOGICAL APPLICATIONS THEREOF

| Al | PPLN. TYPE | SMALL ENTITY | ISSUE FEE DUE | PUBLICATION FEE DUE | PREV. PAID ISSUE FEE | TOTAL FEE(S) DUE | DATE DUE |
|----|--------------|--------------|---------------|---------------------|----------------------|------------------|------------|
| no | nprovisional | NO | \$1440 | \$300 | \$0 | \$1740 | 06/10/2008 |

THE APPLICATION IDENTIFIED ABOVE HAS BEEN EXAMINED AND IS ALLOWED FOR ISSUANCE AS A PATENT. PROSECUTION ON THE MERITS IS CLOSED. THIS NOTICE OF ALLOWANCE IS NOT A GRANT OF PATENT RIGHTS. THIS APPLICATION IS SUBJECT TO WITHDRAWAL FROM ISSUE AT THE INITIATIVE OF THE OFFICE OR UPON PETITION BY THE APPLICANT. SEE 37 CFR 1.313 AND MPEP 1308.

THE ISSUE FEE AND PUBLICATION FEE (IF REQUIRED) MUST BE PAID WITHIN THREE MONTHS FROM THE MAILING DATE OF THIS NOTICE OR THIS APPLICATION SHALL BE REGARDED AS ABANDONED. THIS STATUTORY PERIOD CANNOT BE EXTENDED. SEE 35 U.S.C. 151. THE ISSUE FEE DUE INDICATED ABOVE DOES NOT REFLECT A CREDIT FOR ANY PREVIOUSLY PAID ISSUE FEE IN THIS APPLICATION. IF AN ISSUE FEE HAS PREVIOUSLY BEEN PAID IN THIS APPLICATION (AS SHOWN ABOVE), THE RETURN OF PART B OF THIS FORM WILL BE CONSIDERED A REQUEST TO REAPPLY THE PREVIOUSLY PAID ISSUE FEE TOWARD THE ISSUE FEE NOW DUE.

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IMPORTANT REMINDER: Utility patents issuing on applications filed on or after Dec. 12, 1980 may require payment of maintenance fees. It is patentee's responsibility to ensure timely payment of maintenance fees when due.

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| 10/502,332 | 07/23/2004 | | Marie Malissen | | В | JS-3665-113 | 9194 |
| APPLN, TYPE | : MUTATED GENE CC | ISSUE FEE DUE | TEIN AND THE BIOLO PUBLICATION FEE DUE | PREV. PAID ISSUE | | TOTAL FEE(S) DUE | DATE DUE |
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| EXAM | | ART UNIT | CLASS-SUBCLASS | _ | | | |
| TON, TH | AIAN N ence address or indication | 1632 | 2. For printing on the | | | | |
| CFR 1.363). Change of correspondence address (or Change of Correspondence Address form PTO/SB/122) attached. "Fee Address" indication (or "Fee Address" Indication form PTO/SB/47; Rev 03-02 or more recent) attached. Use of a Customer Number is required. | | | or agents OR, alternat (2) the name of a sing registered attorney or 2 registered patent attorney. | (1) the names of up to 3 registered patent attorneys or agents OR, alternatively, (2) the name of a single firm (having as a member a registered attorney or agent) and the names of up to 2 registered patent attorneys or agents. If no name is listed, no name will be printed. | | | |
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| NIXON & VAN | NIXON & VANDERHYE, PC | | | IAIAN N |
| | BE ROAD, 11TH FLO | OR | ART UNIT | PAPER NUMBER |
| ARLINGTON, VA 22203 | | | 1632 | |
| | | | DATE MAILED: 03/10/2008 | |

Determination of Patent Term Adjustment under 35 U.S.C. 154 (b)

(application filed on or after May 29, 2000)

The Patent Term Adjustment to date is 149 day(s). If the issue fee is paid on the date that is three months after the mailing date of this notice and the patent issues on the Tuesday before the date that is 28 weeks (six and a half months) after the mailing date of this notice, the Patent Term Adjustment will be 149 day(s).

If a Continued Prosecution Application (CPA) was filed in the above-identified application, the filing date that determines Patent Term Adjustment is the filing date of the most recent CPA.

Applicant will be able to obtain more detailed information by accessing the Patent Application Information Retrieval (PAIR) WEB site (http://pair.uspto.gov).

Any questions regarding the Patent Term Extension or Adjustment determination should be directed to the Office of Patent Legal Administration at (571)-272-7702. Questions relating to issue and publication fee payments should be directed to the Customer Service Center of the Office of Patent Publication at 1-(888)-786-0101 (571)-272-4200.

| | Application No. | Applicant(s) |
|---|---|---|
| | 10/502,332 | MALISSEN ET AL. |
| Notice of Allowability | Examiner | Art Unit |
| | Thaian N. Ton | 1632 |
| The MAILING DATE of this communication app All claims being allowable, PROSECUTION ON THE MERITS IS herewith (or previously mailed), a Notice of Allowance (PTOL-85) NOTICE OF ALLOWABILITY IS NOT A GRANT OF PATENT R of the Office or upon petition by the applicant. See 37 CFR 1.313 | (OR REMAINS) CLOSED in () or other appropriate commural (IGHTS). This application is su | this application. If not included ication will be mailed in due course. THIS |
| 1. This communication is responsive to <u>11/26/07</u> . | | |
| 2. 🔀 The allowed claim(s) is/are <u>35, 37, 43, 44, 47, 57-61, 65-6</u> | <u>89</u> . | |
| 3. Acknowledgment is made of a claim for foreign priority u a) All b) Some* c) None of the: 1. Certified copies of the priority documents have 2. Certified copies of the priority documents have 3. Copies of the certified copies of the priority do | e been received. e been received in Application | No |
| International Bureau (PCT Rule 17.2(a)). | ocuments have been received | in this haddrar stage application from the |
| * Certified copies not received: | | |
| Applicant has THREE MONTHS FROM THE "MAILING DATE" noted below. Failure to timely comply will result in ABANDONN THIS THREE-MONTH PERIOD IS NOT EXTENDABLE. 4. A SUBSTITUTE OATH OR DECLARATION must be submin INFORMAL PATENT APPLICATION (PTO-152) which giv | MENT of this application. nitted. Note the attached EXAN | MINER'S AMENDMENT or NOTICE OF |
| 5. CORRECTED DRAWINGS (as "replacement sheets") mus | · , , , - | |
| (a) ☐ including changes required by the Notice of Draftspers | | (PTO-948) attached |
| 1) ☐ hereto or 2) ☐ to Paper No./Mail Date | | (110 540) attached |
| (b) ☐ including changes required by the attached Examiner' Paper No./Mail Date Identifying indicia such as the application number (see 37 CFR 1 | 's Amendment / Comment or i | |
| each sheet. Replacement sheet(s) should be labeled as such in | | |
| DEPOSIT OF and/or INFORMATION about the depo- attached Examiner's comment regarding REQUIREMENT | | |
| | | |
| Attachment(s) 1. ☐ Notice of References Cited (PTO-892) | 5 ☐ Notice of Info | ormal Patent Application |
| Notice of Neterences Cited (FTO-032) Dotice of Draftperson's Patent Drawing Review (PTO-948) | 6. ☐ Interview Sur | , , |
| 3. ☐ Information Disclosure Statements (PTO/SB/08), | Paper No./M | lail Date |
| Paper No./Mail Date4. | | tatement of Reasons for Allowance |
| of Biological Material | 9. | |
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PROPOSED EXAMINER'S AMENDMENT

An examiner's amendment to the record appears below. Should the changes and/or additions be unacceptable to applicant, an amendment may be filed as provided by 37 CFR 1.312. To ensure consideration of such an amendment, it MUST be submitted no later than the payment of the issue fee.

Authorization for this examiner's amendment was given in a telephone interview with B.J. Sadoff on February 26, 2008.

The application has been amended as follows:

1. Replace claim 35 with the following:

A mouse whose genome comprises a mutant LAT gene encoding a mutant mouse LAT protein, wherein the amino acid sequence of said mutant mouse LAT protein comprises a single mutation of a tyrosine located at position 136 of the wild-type mouse LAT protein sequence, wherein the mutation is not a composite mutation of the tyrosine residue at positions 175, 195 and 235 of said wild-type mouse LAT protein sequence, wherein the single mutation of the tyrosine located at position 136 consists of a replacement with a residue preventing the association with SH2 domain of proteins, wherein said mutant LAT gene replaces the endogenous wild-type LAT gene such that the mutant LAT gene is under the control of the regulatory regions of the endogenous LAT gene, and wherein the mouse is homozygous for said mutant mouse LAT gene, and wherein the mouse has a phenotype of exaggerated TH2 cell differentiation.

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2. Cancel claims 38, 46, 48-56, 62-64.

3. In claim 37, replace the term "mutated" with the term – mutant -- in line 2, first word.

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4. Replace claim 47 with the following:

An isolated germ cell or somatic cell from the mouse according to claim 35 or any progeny of said mouse containing the mutant LAT gene.

- 5. In claim 57, delete the term "A" in line 1, first word, and insert the following phrase -- An isolated -- .
- 6. In claim 59, insert the term claim before the term "58" in line 1 of the claim.
- 7. Replace claim 60 with the following:

The mutant mouse gene according to claim 57, wherein the sequence of the mutant mouse LAT gene is SEQ ID NO: 1.

8. Replace claim 61 with the following:

The mutant mouse gene according to claim 57, wherein the mutant sequence encodes a mutant LAT protein comprising the mutated amino acid sequence of exon 7 (SEQ ID NO: 3).

9. Add the following new claims:

65. A mouse whose genome comprises one allele of a mutant mouse LAT gene encoding a mutant mouse LAT protein, wherein the amino acid sequence of said mutant mouse LAT protein comprises a single mutation of a tyrosine located at position 136 of the wild-type mouse LAT protein sequence, wherein the mutation is not a composite mutation of the tyrosine residue at positions 175, 195 and 235 of said wild-type mouse LAT

protein sequence, wherein the single mutation of the tyrosine located at position 136 consists of a replacement with a residue preventing the association with SH2 domain of proteins, wherein said mutant LAT gene replaces the endogenous wild-type LAT gene such that the mutant LAT gene is under the control of the regulatory regions of the endogenous mouse LAT gene, wherein the mouse is a carrier of a null allele of the endogenous LAT gene, and wherein the mouse has a phenotype of exaggerated TH2 cell differentiation.

- 66. The mouse according to claim 65, wherein said mutant LAT gene encodes coding for a mutant LAT protein comprising the mutated amino acid sequence of exon 7 (SEQ ID NO: 3).
- 67. The mouse according to claim 65, wherein said single mutation consists in the replacement of the tyrosine by a phenylalanine (Y-F), an aspartic acid (Y-D) or a glutamic acid (Y-E).
- 68. The mouse according to claim 67, wherein said single mutation consists in the replacement of the tyrosine by a phenylalanine (Y-F)
- 69. An isolated germ cell or somatic cell from the mouse according to claim 65 or any progeny of said mouse containing the mutated LAT gene.

10. Replace the Abstract with the following Abstract:

Abstract

The present invention relates to a mutated gene coding for a mutant LAT protein leading to exaggerated TH2 differentiation. The invention relates to biological structures containing the mutant, particularly non-human LAT gene, mutated animals, plasmids, chromosomal DNAs, embryos comprising the mutated gene, and applications thereof. The invention further relates to screening methods for drugs useful for treatment against asthma and allergy. Otherwise, the invention relates to methods for producing IgE antibodies.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Thaian N. Ton whose telephone number is (571)272-0736. The examiner can normally be reached on 9-5:30 M-F.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Peter Paras can be reached on 571-272-4517. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

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/Thaian N. Ton/ Primary Examiner, Art Unit 1632 Page 6